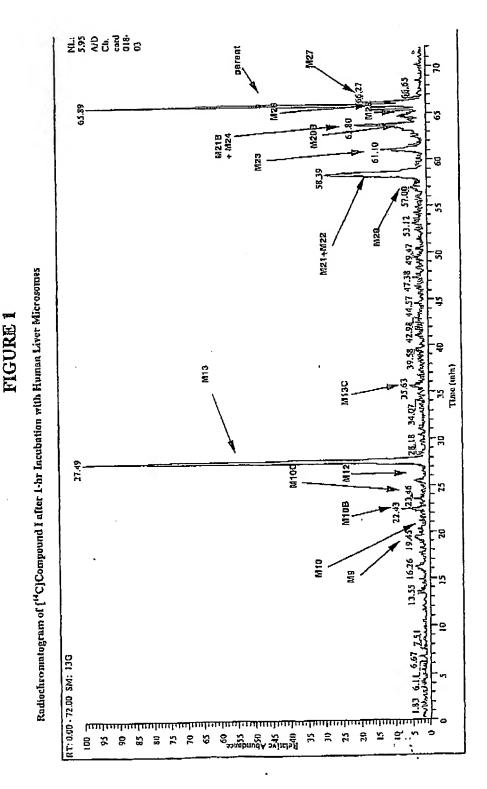
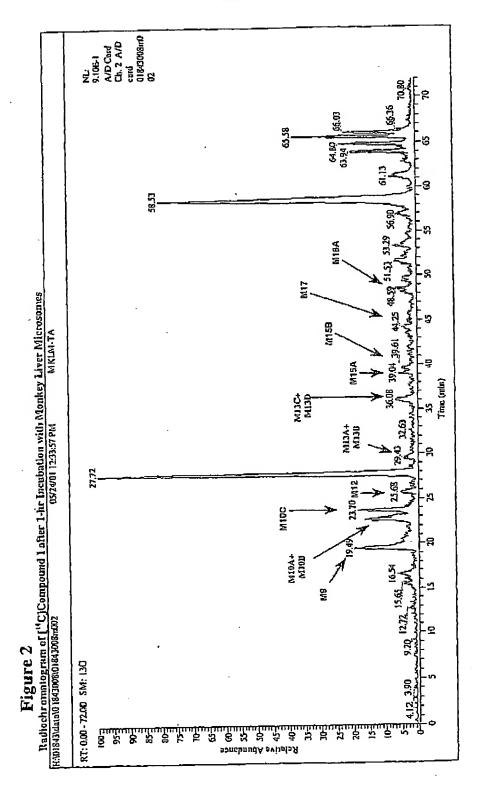
PCT/US2003/034083

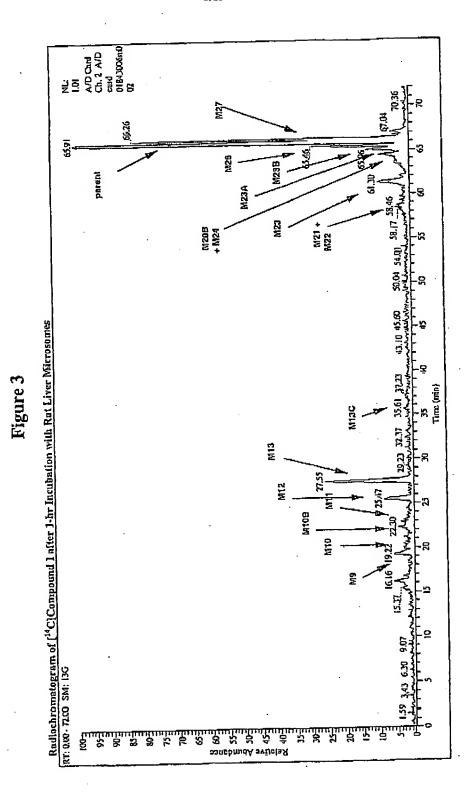
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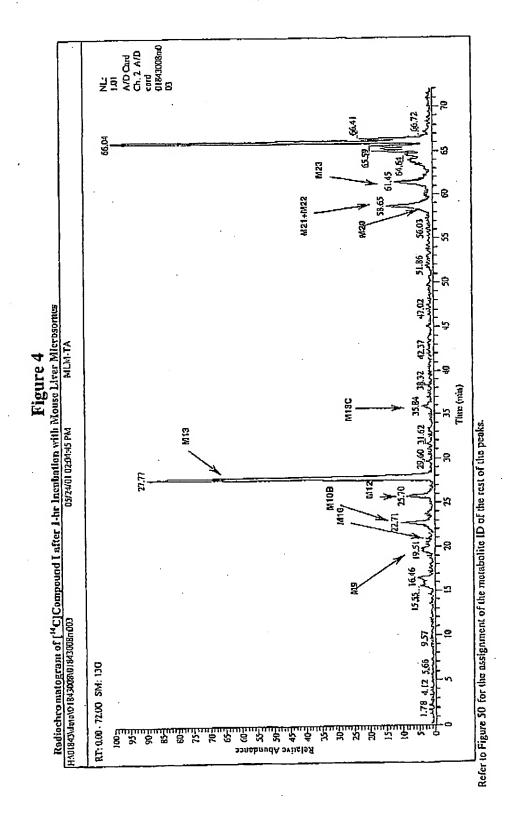
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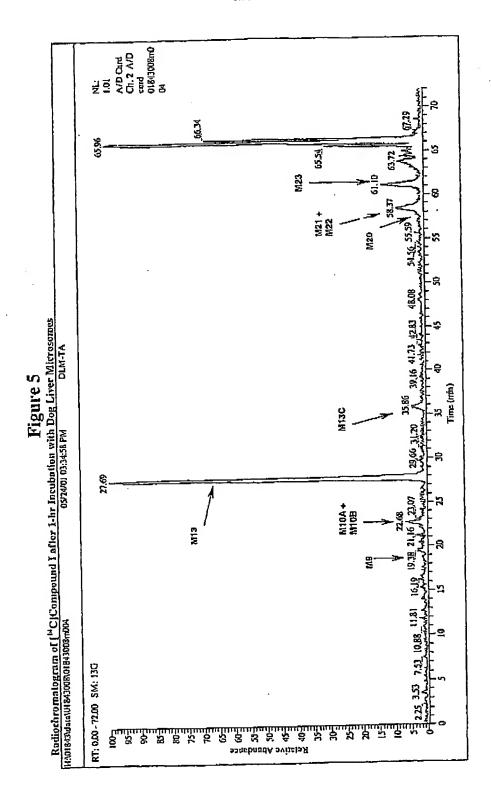
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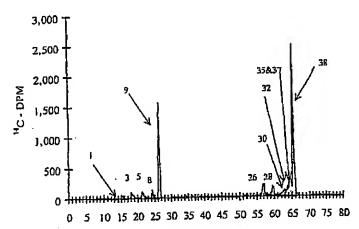


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Figure 6

Reconstructed HPLC Radiochromatogram of [14C]Compound I after 1-hr Incubation with Mouse Liver Microsomes



Time - Minutes

Sample Name: 01843013.002

Start: 11-May-01 LSC Reference: LSC977: 101-1

Method: 01843 ID: X977:49A (MLM-TA)

Column: Acc 5 C18, 4.6 x 250 mm, 5µm, #A3538; Guard: RP-8, 4.6 x 30 mm, 5µm

Column Temperature: 30°C

HPLC System T Detector: UV at 254 nm; 14C: Raytest

Gradiere 100% A, 3 rais 10 40% B, 30 min to 60% B (15 min), 10 min to 65% B, 5 min to 100% B (9 min)
A: 0.01 M NH4OAE (nH=3 w/ HCOOH) B: Methanol Flow Rate: 1.0 ml/min

^N C Data Region No.	Werapolite	Slam	Stop	R,	dpm_	%ROL_	%Total
.טיו ווטרבים	11,220,001114	15.25	15.50	15.50	113	1.09%	0.94%
t	M9 + M10	18.00	19.25	18.00	319	3.13%	2.70%
	м10B	21.00	23.75	21.25	266	261%	3.25%
	M12	24.00	24.50	24.25	229	2.25%	1.94%
	M13	26.00	27.00	26.35	2706	26.58%	23.91%
	M20 + M21 + M22	56.25	57.75	57.00	663	6.51%	5.61%
5	M23	59.25	60.25	59.50	451	4.43%	3.B2%
3	M20B + M21B + M24	61.75	63,25	63.00	410	4.03%	3.47%
3	M23A	63.50	64,00	63.75	364	3.57%	3.08%
2	M23B+M23C	64.25	64.50	64.50	551	5.41%	4.66%
5		64.75	65,00	64.75	395	3.89%	3.34%
7	M23D+M26	65.25	66.25	65.25	3717	36.51%	31.47%
В	Parent DPM Recovered:	11813		PM in ROI:	10182	\$00.00°	86.199
		12465					

DPM Injected; 12465 % Recovery: 94.77

Note: 1) Those peaks which accounted for >0.5% of the total were considered as ROL

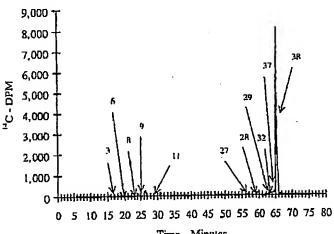
2) M13C and M27 accounted for >0.5% of the total and considered as detacled in a trace amount.

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Figure 7

Reconstructed HPLC Radiochromatogram of [14C]Compound I after 1-hr Incubation with Rat Liver Microsomes



Time - Minutes

Sample Name: 01943011.D03

Start: 9-May-01 LSC Reference: LSC977: 52-83 Method: 01843

ID: X977:48A (RLM-TA)

Column: Ace 5 Ci 8, 4.6 x 250 mm, 5µm, #A3538; Goard: RP-8, 4.6 x 30 mm, 5µm

Column Temperature: 30°C HPLC System T

Detector: UV at 254 nm: "C: Rayrest

Gradient: 100% A, 3 mis to 40% B, 30 min to 60% B (15 min), 10 min to 65% B, 5 min to 100% B (9 min) A: 0.01 M NH₄OAc (pH=3 w/HCOOH) B: Methanol Flow Rate: 1.0 mL/min

¹⁴ C Data Region No.	Membrite	Start	Stop	· R,	dpm	%RO1	%Total
3	M9+MID	18.25	19.25	18.25	183	1.29%	1.16%
6 .	M10B+M11	21.00	22.00 .	21.25	151	1.06%	0.96%
8	M12	24.25	24.50	24.25	174	1.23%	1.11%
9	M13	26.25	26.75	26.50	554	3.91%	3.52%
11	14415	30.75	31.00	30.75	173	1.22%	1.10%
27	M21 + M22	56.50	57,25	56.75	159	1.12%	J.01%
28	M23	59.00	60.25	59.50	362	2.55%	2.30%
29	M20B + M24	62.00	63.25	62.75	212	1.49%	1.35%
32	M23A	63.50	64.00	63.75	234	1.65%	1.49%
37	M23B + M26	64.25	65.00	64.75	572	4.03%	3.63%
_ :	porent	65.25	66.50	65.25	11411	BD.44%	72.48%
38	DPM Recovered:	15744		PM in ROt	14185	99.99%	90.11%
	DPM Injected:	15460					
	% Recovery:	101.84					

Note: 1) Those peaks which accounted for >0.5% of the total were considered as ROL

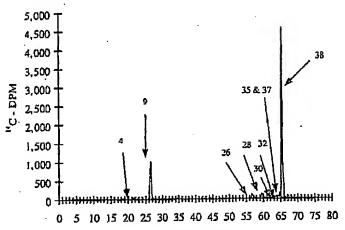
2) M13C and M27 accounted for <0.5% of the total and considered as desected in a trace amount.

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Figure 8

Reconstructed HPLC Radiochromatogram of [14C]Compound I after 1-hr Incubation with Dog Liver Microsomus



Time - Minutes

Sample Name: 01843013.003

Siari: 11-May-01 LSC Reference: LSC977: 114-115

Method: 01843

ID: X977:498 (DLM-TA)

Column: Acc 5 C18, 4.6 x 250 mm, 5µm, #A3538; Guard; RP-8, 4.6 x 30 mm, 5µm

HPLC System T Column Temperature: 30°C

% Recovery: 93.99

Desector: UV at 254 nm; IAC: Raytest

Gradjent: 100% A. 3 min to 40% B, 30 rain to 60% B (15 min), 10 min to 65% B, 5 min to 100% B (9 min)
A: 0.01 M NH₄OAc (pH=3 w/ HCOOH) B: Methanal Flow Rate: 1.0 mL/min

¹⁴ C Data Region No.	Matabolite	Slart	Stop	R,	dpm	50 No.	%Total
4	MIOA + MIOB	21,25	21.50	21.50	68	0.72%	0.64%
9	M13	26,25	27.00	26.50	1807	19.15%	17.10%
76	M20 + M21 + M22	56.50	57.25	56.75	255	2.70%	2.41%
28	M23	59,25	60.00	59.50	321	3,40%	3,04%
30	M208 + M218 + M24	62.00	63.25	62.75	256	2.71%	2.42%
32	M23A	63.50	64.00	63.75	195	2.07%	1.85%
35	M23B + M23C + M25	64.25	64.50	64.25	103	1.08%	0.97%
37	M23D + M26	64.75	65.00	64.75	235	2.49%	2.22%
38	pareni	65.25	66.50	ნ5.25	6198	65.68%	58.65%
,,,,	DPM Recovered:	1056B		M in ROL	9437	100.00%	89.30 %
	DPM Injected:	11244					

Note: 1) Those peaks which accounted for >0.5% of the total were considered as ROI.

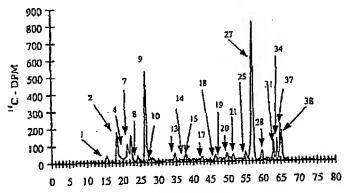
2) M9, M13C and M27 accounted for <0.5% of the total and considered as detected in a trace amount.

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Figure 9

Reconstructed HPLC Radiochromatogram of [14C]Compound I after 1-hr Incubation with Monkey Liver Microsomes



Time - Minutes

Sample Name: 01643013.D04

% Recovery: 101.65

Method: 01843 ID: X977:49C (MKLM-TA)

10-YeM-11 :1rai2 LSC Reference: LSC977: 105-10:

• .							•
"C Data Region No.	Mecabolica	Start	Ştop	R.	dpm	%ROI	%Total
ICEBION 140-		15.25	15.75	15.50	92	1.37%	1,125
	M9	18.00	20.50	18.25	59 I	8.80%	7.13%
	M10A + M10B	20.75	22.00	21,25	367	5.46%	4.46%
4	MIDC	22.25	22.75	22,25	278	4.14%	3,38%
1	M12	24,25	24,50	24.25	. 61	0.91%	D.7456
3	M13	26.00	27.00	25.25	1107	16.48%	13.45%
)	BEIM + AELM	27,75	28.25	28.25	73	1.09%	0.89%
10		34.50	34.75	34.75	88	1.31%	1.07%
13	MISA	37.50	38.00	37.75	95	1,41%	1.15%
14	MISB	38.25	38.50	38.25	56	0.63%	0.68%
15	M17	42.50	42.75	42.75	57	0.85%	\$26.0
17	MIBA	46.25	46.75	46_50	88	1.31%	1.07%
18		47.50	47.75	47.75	41	0,61%	0.50%
19	MIBB	49.25	30.50	49.75	189	2.81%	2.30%
20	M18C + M19	51.00	51.75	51.50	119	1.77%	1.45%
21	M19A + M19B	54.75	55.50	55.00	156	2.32%	1.90%
25	M20	56.00	57.75	57.00	2176	32,39%	26,44%
27	M21 + M22	59.50	60.00	59.75	137	2,04%	1.66%
28	M23	62.75	63.50	63.00	208	3.10%	2,53%
31	M21B + M24		64,25	64.00	191	2.84%	2,32%
34	M25	63.75		64,75	307	4.57%	3.73%
37	M23D + M26	64.75	65.00	65.25	242	3.60%	2.94%
38	Porent	65.25	65.75	PM in ROE	6719	100.01%	81.65%
	DPM Recovered:		ט	rw in KOE		100.01 20	0,005,12
	DPM Injected	8097			•		

Note: 1) Those peaks which accounted for >0.5% of the total were considered as ROL

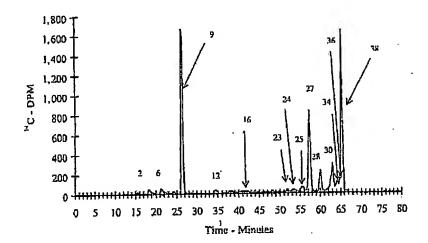
²⁾ M27, and Regions 16, 23, and 24 accounted for <0.5% of the total and considered as detected in a trace ompubit

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Figure 10

Reconstructed HPLC Radiochromatogram of [14C]Compound I after 1-hr Incobation with Human Liver Microsomes



Sample Name: 01843010-D02 Method: 01843

% Recovery: 102.18

Scan: 6-May-01 LSC Reference: LSC977: 88, 90-91

ID: X977:49D (HLM-TA)

Column: Att 5 C18. 4.6 x 250 mm. 5µm. #A3538; Guard: RP-8, 4.6 x 30 mm, \$µm Column Temperature: 30°C

HPLC System T

Detector: UV of 254 hm; 14C: Raylest

Gradient 100% A, 3 min to 40% B, 30 min to 60% B (15 min), 10 min to 65% B, 5 min to 100% B (15 min)
A: 0,01 M NH₂OAc (pH=3 w/ HCOOH) B: Methanol Flow Rate: 1.0 mL/min

"C Data Region No.	Memballic	Start	Stop	R,	₫pm	%RO)	Total
2	M9	18.25	18,75	18,25	114	0.97%	0.85%
2 6	MIOB	21.25	21.75	21.25	129	1.10%	0.95%
9	MI3	26.00	27,00	26.25	3571	30,53%	26.66%
12	MI3C	34.25	34.75	34.50	99	0.85%	0.74%
16		41,25	42.25	42,00	112	0.96%	0.84%
23		51.75	52,25	51.75	BB	0.75%	0.66%
24		52.75	\$4.25	\$3.50	168	1.44%	1,25%
24 25	M20	55.00	56.25	55.50	251	2.15%	1.87%
27	M21 + M22	56.50	58,25	57.25	2122	IR.14%	15.84%
20 28	MZ3	59.50	60.75	60:00	591	5.05%	4,41%
30	M208 + M218 + M24	61.25	63.50	63.00	1044	8.91%	7.79%
	M25	63.75	64.50	64.00	390	3.33%	2.91%
34 36	M26	64.75	64.75	64.75	145	1.24%	3.08%
36	paren)	65.00	66.00	65.25	2871	24.55%_	21,43%
38	DPM Recoverests	13396		PM in ROL	11695	99,99%	B7.29%
	DPM Injected:	13110					

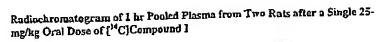
Note: 1) Those peaks which accounted for >0.5% of the total were considered as ROL.

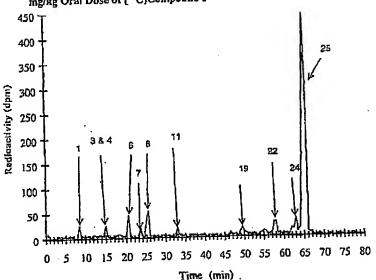
²⁾ Each of M10, M10C, M12, and M27 accurated for <0.5% of the total and considered as descented in trace amounts.

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FIGURE 11





Sample Name: 01842009.D02

Method: 01842

ID: X966:29E (Pooled Plasma, 1 hr)

Start: 3-Apr-01 LSC Reference: LSC966:84-87

Column: Ace 5 C18, 4.6 x 250 mm, 5 μm, 6A3538, Guard; RP-8, 4.6 x 30 mm, 5 μm

HPLC System T. Column temperature: 30 °C

Deterior: UV at 254 nm; HC: Roylest

Gradient: 100% A, 3 min to 40% B, 30 min to 60% B (15 min), 10 min to 55% B, 5 min to 100% B (15 min)

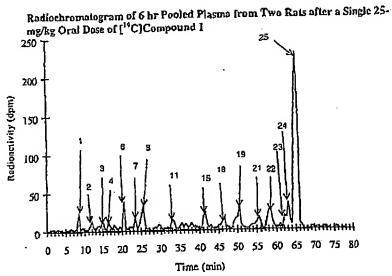
A: 0.01 M NH,OAc (pH=3 art HCOOH) B: Methanol; Flow Rate: 1.0 mL/min

Region No.	"C Data Name	Start	Stop	R,	dpm	%ROI	%Total
		8.5	· 8.5	8.5	25	1.64%	1.35%
	M3		15.0	15.0	25	1.84%	1.35%
& 4	M7 + M8 + M8A	15.0	20.5	20.5	46	3.39%	2.48%
5	M10	20.5		23.5	22	1.62%	4.1 9%
•	M12	23.5	23.5	25.5	96	7.07%	5.18%
}	M13	25.0	25.5	-	20	1.47%	1.08%
1	M14	0.KE	33.0	33.0	19	1.40%	1.03%
9		49,5	49.5	49.5	57	4.20%	3.08%
22	M23	57.5	58.0	<i>57.5</i>		4.94%	3.62%
24		62.0	63.0	63.0	67	•	52.92%
<u>2</u> 5 _	trong	64.0	65,5	65.0	980	72,72%	73.28%
	DPM Recovered: DPM Injected: Recovery:	1852 1770 104.63		DPM in ROE Note: any fraction as background a	1357 on containing nd not includ	99.99% less than 15 dpm ed in the integral	was conside

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FIGURE 12



Sample Name: 01842009.D03

Method: 01842

% Recovery: 112.57

ID: X966:29F (Pooled Plasma, 6 hr)

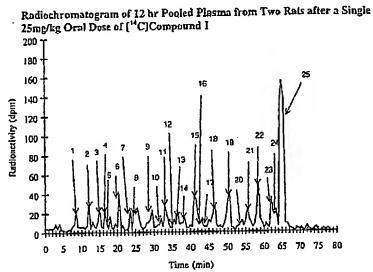
Sun: 9-Apr-01 LSC Reference: LSC966:89-92

Regian No.	Name	Stort	Stop	R _r	dpm	%ROI	%Total
	M3	8.5	. 8.5	8.5	24	2.33%	1.57%
l 2	1/12	12.0	12.D	12,0	14	1.36%	0.91%
	1.47	15.0	15.0	15.0	16	1.55%	1.05%
}	M7	16.5	16.5	16.5	12	1.16%	0.78%
	M8 + M8A	20.5	20.5	20.5	38	3.69%	2.48%
•	MID		23.5	23.5	17	1.65%	1.11%
7	M12	23.5	25.5	25.5	52	5.04%	3.40%
8	M13	25.0	33.5	33.0	28	2.72%	1.83%
1 (M14	33.0		41.5	43	4,17%	2.81%
15	M16	41.0	41.5	46.5	17	1.65%	1.11%
18	M19	46.5	46.5	50. <i>5</i>	62	6.01%	4.05%
19		49.5	50.5		17	1.65%	1.11%
21	M21 or M22	55.5	55.5	55.5 50.0	64	6.21%	4.18%
22	M23	57.5	58.5	58.0	34	3.30%	2,22%
23	M24	62.0	62.5	62.0	-	4.75 %	3.20%
24		63.0	63.5	63.0	49		35.53%
25	parent	64.5	65.5	65.0	544	52.76%	67.34%
	DPM Recovered:	153)		DPM in ROI:	1031	100.00%	
	DPM Injected:	1360		Note: any frac	tion contain	ing less than 12	dpm was coas
	% Recovery:	112.57		as background	and not ind	cluded in the into	egration.

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FIGURE 13



Sample Name: 01842009.D04 Method: 01842

• ID: X966:29G (Pooled Plasma, 12 hr)

Start: 9-Apr-01 LSC Reference: LSC966:93-96

	"C Data Name	Sinri	Sipp	R,	dpm	% ROI	%Total
			0.5	B.5	21	1.59%	1.11%
3	M3	B.5	B.5	12.0	28	2.12%	1,48%
2 3		12.0	12.0	15.0	35	2.65%	1.85%
	M7	14.5	15.0	16.5	21	1.59%	1,11%
4	MB + MBA	16.5	16,5 18.0	1B.O	16	1.21%	0.84%
5	M9	18.0		20.5	40	3.03%	2.11%
6	MIO	20.5	20,5 23.5	20.5 23.5	23	1.74%	1.21%
7	M12	23.5		25.5	83	6.28%	4.38%
6 7 8 9	M13	24,5	26.0	29.5	36	2.73%	1.90%
		29.0	29.5	32.D	14	1.06%	0.74%
10		32.0	32.0	33.0	45	3.41%	2.37%
11	M14	33.0	33.5	35.0 35.5	13	0.98%	0.69%
12	M15	35.5	35. 5	36.5	22	1.67%	1.16%
13		36.5	36.5	38.5	14	1.06%	0.74%
14		38.5	38,5		87	6.59%	4.59%
15	M16	41.0	42.0	41,0	12	0.91%	0.63%
16		43.0	43.0	43.0	12	0.91%	0.63%
17	M18	45.0	45.0	45.0	46	3.48%	2.42%
18	M19	46.D	46.5	46.5		6.66%	4.64%
19		49.5	50.5	50.5	88 12	0.007	Q.63%
20		54.D	54.0	54.0		3,79%	2,64%
21	M21 or M22	55.0	56.0	55.5	50		4.38%
22	M23	5B.Ô	59.0	5B.5	83	6.28%	2,95%
23	M24	62.0	62.5	62.D	56	4.24%	
24		63.0	63.5	63.0	40	3.03%	2.11%
25	parent	64.5	65.5	65.0	424	32.10%	22.35%
	DPM Recovered:	1897		DPM in ROI:	1321	100.02%	69.66%
		1 5770		Material Core	tion contain	ing loce than 12	dom was co

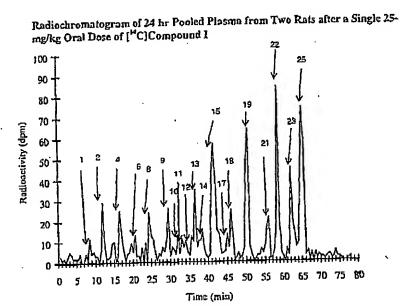
DPM Recovered: 1897
DPM Injected: 1570
Since Recovery: 120.83

Note: any freetian containing less than 12 dpm was considered as background and not included in the integration.

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FIGURE 14



Sample Name: 01842009.D05

Method: 01842

DPM Injected:

% Recovery: 117.62

1D: X966:29H (Pooled Plasma, 24 hr)

\$18ft: 9-Apr-01 LSC Reference: LSC966:97-101

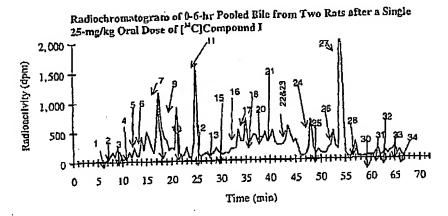
Region No.	¹⁴ C Data Name	Start	Stop	R,	dpm	%ROI_	%Total
110.		8.5	B.5	B.5	11	1.10%	0.77%
1	М3		12.0	12.0	29	2.89%	2.02%
2		12.0	17.0	16.5	37	3.69%	2.58 %
4	M8 + M8A	16.5	-	20.5	13	1.30%	0.91%
6	м10	20.5	20.5	24.5	48	4.79%	3.34%
8	M13	24.5	25.5	29.5	37	3.69%	2,58%
9		29.0	29.5	32.0	13	1.30%	0.91%
10		32.0	32.0	33.0	ii	1.10%	0.77%
11	M14	33.0	33.0	35.5	12	1.20%	0.84%
12	M15	35. <i>5</i>	35.5	35.5 35.5	35	3.49%	2.44%
13		36.5	36.5	38. 5	14	1.40%	0.98%
14		38.5	38.5	41.5	166	16.57%	11.57%
15	M16	41.0	43.0	45.0	13	1.30%	0.91%
17	M18	45.0	45.0	45.0 46.0	38	3,79%	2.65%
18	M19	46.0	46.5	50.5	113	11.28%	7.87%
19		50.0	50.5	56.0	38	3.79%	2.65%
21	M21 or M22	55.5	56.0	58.5	119	11.88%	8.29%
22	M23	58.0	59.0	62.0	80	7.98%	5.57%
23	M24	62.0	63.0		175	17.47%	12,20%
25	parent	64.5	65.5	65.0		100.01%	69.85%
	DPM Recovered:	1435		DPM in ROI:	1002	100.0170	- 40.00

Note: any fraction containing less than 11 dpm was considered as background and not included in the integration.

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FIGURE 15



Sample Name: 01842006.D02 Method: 01842 ID: Pooled Bile 0-6 for (X966:17A) Store 30-Mar-01 LSC Reference: X966:37-40

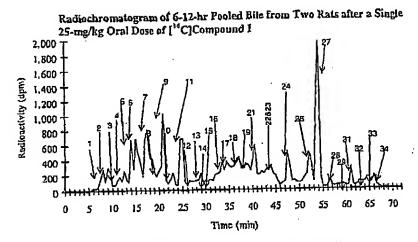
Region	nC Dam		_		٠	%ROI	firoT#	% Dasc
No.	Name	Sun	Sipp	R,	dpm	1.16%	1.16%	0.16%
1	M1 + M2	7.0	8.5	8.0	367	1.50%	1.49%	0.20%
2 3	M4 + M5	9.0	10.5	9.0	473	0.94%	0.93%	0.12%
3		11.0	12.0	11,5	256		1.37%	0.18%
4		12.5	13.5	125	436	1.38%	1.95%	0.26%
5	M5	14.0	14.5	14.0	617	1.95%	4.11%	0.55%
6	M7 + MB + MBA	15.0	16,5	15.0	1303	4.12%	9.58%	1.28%
7	M9	17.0	19.0	17.5	3037	9.60%	2025	0.27%
В	MID	19.5	20.5	20.0	641	203%	202 7 3.55%	0.48%
9	MH	21.0	22-0	21.0	1125	3.56%		0.20%
10	MI2	22.5	24.D	23.0	475	1.50%	1.50%	0.90%
11	M13	24.5	25.0	25.0	2138	6,76%	6.74%	0.14%
12		26.5	27.5	27.0	334	1.06%	1.05%	
13		28.0	30.0	29.0	730	2.31%	2.30%	0.31%
15		30.5	31.5	31.0	365	1.15%	1.15%	0.15%
16	M14	32.D	34.0	33.5	1425	4.51%	4.49%	0.60%
17	M15	34.5	35.5	35.0	1282	4.055	4.04%	0.54%
18		36.0	37.5	36.5	1532	4.84%	4.83%	0.65%
20		38.0	39.5	39.0	1545	4.89%	4.87%	0.65%
21	M)6	40.0	41.5	40.5	1447	4.58%	4.56%	0.61%
22 & 23	M17+M18	42.0	46.5	43.5	2831	8.95%	₽. 93₩	1.20%
24 24	M19	47.0	49.0	48.0	1224	3.87%	3.86%	0.52%
24 25	1.675	49.5	50.5	49.5	390	1,23%	1.23%	0.16%
25 25	M20	51.0	53.5	52.5	1412	4.46%	4,45%	0.60%
	M22	54.D	56.0	54.0	4034	12.76%	12.725	1.70%
27	M23	56.5	59.0	57.0	587	1,66%	1.85%	0.25%
28	دعيهم	59.5	60.0	59.5	106.	0.34%	0.33 7	0.1045
30	3.27.4	60.5	62.0	61.5	37D	1.17%	1.17%	0.1 6%
31	M24	62.5	63.5	63,0	302	0.95%	0.95%	0.13%
32		64.D	65.5	65.0	586	1.85%	1.85%	0.25%
33	patent	66.0	67,0	66.0	214	0.69%	0.67%	0.09%
34				M in ROI:	31624	100.01%	99.70%	13.35%
	DPM Recovered:	21113	יוע	at at vol-	21047	,		

DPM Injected: 32752 % Recovery: 96.83

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FIGURE 16



Sample Name: 01842006.D03 Method: 01842 10: Pouled Bile 6-12 lg (X966:170) Stort: 30-Mar-01 LSC Reference: LSC066:41-44

Regios	"C Date		•					
No.	Name	Sum	Stop	R,	dpm		. STotal	% Drac 0.15%
	MI + M2	6.5	B.5	6.0	612	2.005	£20.5	
2	M4 - M5	90	10,5	4,0	67B	2.32G	230%	0.15%
3 .		11.D	12.0	11.5	442	1.51%	1.50%	6.115
4		12,5	13.5	125	511	1.75%	1.74%	0.12%
5	M6	14.0	14.5	D.AI	947	3,24%	3.72%	0.23%
6	M7 + MB + MBA	15.D	16.5	15.D	1704	5.83%	5.79%	0.41%
7	M9	17.0	19,0	17.5	2596	E.£7‰	8.B2%	0.62%
B	MID	19.5	20.5	20.0	745	255%	153%	0.18%
>	MII	21.0	22.0	21.0	1231	4,21%	4.18%	0.295
ID.	M12	22.5	24.0	23.0	504	1.72%	1.71%	0.12%
11	M13	24.5	26.0	25.0	1546	\$2 \$ %	5.35%	0.37%
13		26.5	235	26.5	351	1.70%	L19%	ODBT
(3		28,0	29.5	29.0	562	1,925	1.91%	0.13%
14		30.0	30.5	30.D	233	O'BO#	0.79%	0.06%
13	•	31.0	31.5	31.0	360	0.89%	0.86%	CL06%
16	M14	32,0	34.0	23.5	1282	4.38%	4.36%	0,31%
17	M15	34.5	35.5	34.5	290	3.38%	3.36%	0.24%
18	61.5	36.0	37.5	37.0	1392	4.76%	4.73%	0.33%
19		38.0	39.5	38.5	1197	4.09%	4.075	0.289
2)	M16	40.0	41.5	40.5	1422	4.86%	4.83%	0.34%
22 & Z3	M17+M18	12.0	46.D	44 D	1856	6.35%	6.31%	0.44%
24	MIP	46,5	50.0	47.5	1403	4.EO/2	4.77%	0.33%
24 25	M20	50.5	.53.0	52.0	1372	4,69%	4,65%	0.73%
27 27	MZZ	53.5	35.5	54.0	3479	11.89%	11.825	CB.O
-	M23	36.0	57.5	56.5	260	0.89%	0.68%	0.069
28	1*84.3	58.0	60.0	59.0	259	0.89%	₹88,0	0.06%
29	M24	60.5	67.0	61,0	412	1,41%	1.40%	0.107
31	1474	62.5	63.5	63.0	273	0.935	0,93%	0.075
32		60.0	65.5	64.5	475	1.62	1.61%	0.119
33	barent	66.0	67.0	6G.D	257_	0.68%	0.87%	0.069
34	5514M . 15			PM in ROI:	79253	100.00%	99.37%	6.98%
	DPM Recovered		u	TH III VOL		,		
	DPM Injected:	30078						

S Recovery: 96.88

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FIGURE 17

Radiochromatogram of 12-24-hr Pooled Bile from Two Rats after a Single 25-mg/kg Oral Dose of [$^{14}{\rm C}$]Compound I 2,000 1,800 1,600 Radionalvity (dpm) 1,400 1.200 1,000 800 600 400 65 70 бO žΟ 55

30 35 40 45

Time (min)

Sample Name: 01842006.D04 Method: 01842 ID: Pooled Bile 12-24 hr (X966:17C)

10 15 20 25

Start: 30-Mur-01 LSC Reference: LSC966:45-48

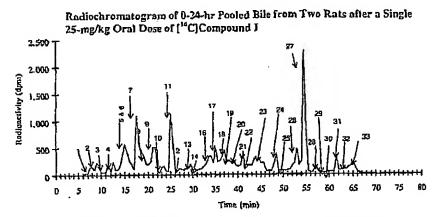
Region	¹⁴ C Data						 .	% Dose
No.	Name	Staff	Stop	R,	क्रमा	5ROI	%Total	0.U9%
1	M1+M2	6.5	8,5	B.D	434	€83.1	1.68% 1.80%	0.09%
	744 ÷ 145	9.0	10.0	9.0	467	1.61%		0.10 ₹
3		1D.5	12.0	11.5	394	1.535	1.52% 1.45%	D.08%
2 3 4 5 6 7		12.5	13.5	12.5	375	1.45%	2.24%	0.12%
Ġ	M6	14.0	14.5	14.0	581	2,25%		0.1279
6	M7 + M8 + M8A	15.0	16.5	15.0	1273	4.94%	4.92% 8.61%	0.46%
ř	M9	17.0	19.D	17.5	7229	8.65%	2.45%	0.13%
Ė	M10	19.5	20.5	20.5	633	2.46%	7.315	0.137
9	M11	21.0	22,0	21.0	B56	3.72%	1.56%	0.08%
10	MI2	22.5	24. D	23.D	404	1.57%	2.62 7	0.567
ii	M13	24.5	26.0	25.0	1454	5.64%	1.15%	0.069
12		26.5	27.5	27.0	297	1.15%	1.97%	0.00%
13		28.0	20.5	29.D	510	1.98%		0.017
14		30.0	30,5	30.0	207	0.80%	0.80%	
15		31.0	31.5	31.0	727 .	0.86%	0.B6%	0.059
15	M14	32.0	34.0	33,5	1107	4,30%	4.28%	0.237
17	M15	34.5	35.5	34.5	942	3.65%	3.64%	0.207
18		36.0	39.0	36.5	1642	6.37%	6.34%	0.347
19		38.5	39 .5	38.5	B64	3.35%	3.34%	Ø.189
21	M16	40.0	41.0	40.0	1093	4.24%	4, 22%	0.335
22	M17	41.5	42.5	41.5	618	2.40%	2.39%	0.139
23	MIB	43.0	46.D	43.5	1328	5.15%	5.13%	0.285
24	M19	46.5	48.5	47.5	1087	4.22%	4,20%	0.239
25	,,,,-	49.0	49.5	49.0	208	0.81%	708.0	0.049
26	M20	50.0	£3.D	52.0	1483	5.75%	5.73%	0.319
27	M22	53.5	55.5	53.5	3244	12.59%	1253%	0.689
	M23	\$6.0	57.5	56.5	255	0.99%	0. 99 %	0.05
28	11147	58.0	58.5	58.0	93	0.36%	ቤ 36%	0.024
29		59.0	60.0	59.0	156	0.61%	0.60%	0.03
30	M24	60.5	62.0	0,10	338	1.31%	1.31%	0.07
31	F124	62.5	64.0	63.5	348	1.35%	1.34%	0.07
32		64.5	65.5	64.5	359	1.39%	1.19%	0.08
33	parent		67.0	66.0	273	1.06%	1,05%	0.05
34		66.0		PM in ROI:	25774	99.99%	99,58%	5,385
	DPM Recovered:	25882	D	rm is KUI:	2114	77.2,20	3,000	
	OPM Injected:	26360						

% Recovery: 98.19

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FIGURE 18



Sample Name: 01842009.003 Method: 01842 ID: X966:288 (Pooled Bile, 0-34 hr)

Start: 3-Apr-01 LSC Reference: LSC966:80-83

Regico	[™] C Dain							
No.	Name	Ştan	Stop	R ₂	dpm	%ROI	[2007ल	% Dose
7	M1 + M2	6.5	8.5	6.0	447	1.68%	1.58%	0,43%
2	M4 + M5	9.0	2.01	9.0	502	1.91%	1.90%	0.49%
3		11.0	12.0	11.5	329	1.25%	1.25%	0,32%
4		12.5	13.5	17.5	414	1.58%	1.57%	0.419
S & 6	M6 + M7 + M8 +M8A	14.0	17.0	t5.0	2111	ደውም	5.00%	2.069
7	MD	17.5	19.5	17.5	2671	10.17%	10.125	2,619
8	MID	30.0	20.5	20.5	432	1.64%	1.64%	0.429
9	MIL	21.0	22.5	21.0	1141	4.34%	4.32%.	1,115
10	M12	23.0	24.5	23.5	439	1.67%	1.66%	0.439
11	M13	25.0	26.5	25.0	1853	7.05%	7.02%	1.819
÷2		27.0	ZR.0	27.0	312	1.19%	1.18%	0,309
13		28.5	30.0	29.5	536	2.04%	2,03%	0.529
14		30.5	31.0	30.5	182	0.69%	0.69%	0.189
16	M14	31,5	34.5	34.0	1477	5.62%	5.60%	1,44
17	M15	35.0	35.5	35.0	748	2.85%	2.84%	0.73
1 F	11172	36.0	37.5	37.0	1269	4.83%	4.81%	1.24
19		38.0	38.5	38.0	501	1.91%	1.90%	0.49
20		39.0	40.0	39.0	612	2.33%	2.32%	0.60
20 21	міб	40.5	41.5	41.0	693	2.645	2.63%	0.69
21 22	M17	42.0	43.0	42.0	537	2.04%	2.04%	0.53
23	MIB	43.5	46.5	43.5	1500	5.71%	5.69%	1:47
	MI9	47.0	49.0	48.5	757	2.88%	2,87%	0.74
24 25	MIZ	49.5	50.5	50.0	198	D.75%	0.75%	0.19
-	M20	\$1.0	53.5	57.0	1174	4.47%	4.45%	1,15
26	M22	54.0	56.0	54.5	3444	13,11%	13.05%	3.37
27		56.5	58.0	57.0	307	1.17%	1.16%	0.30
28	M23	58.5	59.0	58.5	61	0.23%	0.23%	0.06
29		59.5	60.5	60.0	157	0.60%	0.60%	0.15
30		_	62.5	61.5	527	2.01%	2.00%	0.52
31	M24	61.0		63.0	223	0.85%	0.85%	0.22
32		63.0	63.5	65.0	720	2.74%	2.73%	0.70
33	parent	64.0	66,5				99.58%	25.67
	DPM Recovered:	26382		DPM in ROI:	26269	99.99%	מהסניהג	23.07

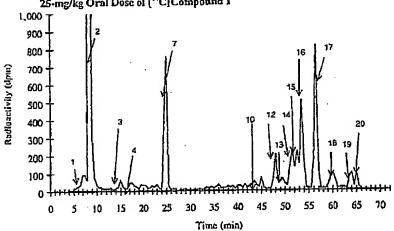
DPM Injected: 24582 % Recovery: 107.32 WQ 2004/037211

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FIGURE 19

Radiochromatogram of 0-12-hr Pooled Urine from Two Rats after a Single 25-mg/kg Oral Dose of [14 C]Compound I



Sample Name: 01842006.005 Method: 01842

(D: Pooled Urine 0-12hr (X966:17D)

Stort: 30-Mar-01 LSC Reference: LSC966:49-52

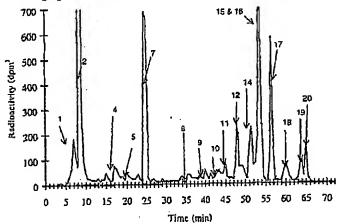
Region	IIC Data	5	Pi	ኤ ,	dpm	%ROI	%Тош)	% Dose
No.	Name	Start	Stop			2.24%	2.05%	0.06%
F .		6.0	B.0	7.0	3)0			
2	M3	8.5	10.0	8.5	8223	59.55%	54.41%	1.47%
3.	M7 + M8 + M8A	15.0	15.5	15.0	90	D.65%	0.60%	0.02%
4	M9)7.0	18.5	17,5	151	1.09%	1.00%	0.03%
7	M13	24.5	25.5	25.0	1181	₿.55 %	7.81%	0.21%
10	M17	42.5	44.0	43,5	151	1.09%	1.00%	0.03%
12	M19	47.5	49.0	4B.0	3B1	2.76%	2.52%	0.07%
13	,	49.5	50.5	49.5	135	0.98%	0.89%	0.02%
14		51.0	52.0	51.5	499	3.61%	3.30%	0.09%
15	M20	52.5	53.0	52.5	371	2.69%	2.45%	0.07%
16	M21	53.5	54.0	53.5	660	4.7E%	4.37%	0.12%
	M23	56.0	` 57.0	56.5	1113	8.06%	7.36%	0.20%
17	M24	59.0	60.5	60.0	267	1.93%	1.77%	0.05%
IB	NIZ4	63.5	64.0	64.0	174	1.26%	1.15%	0.03%
19			65.0	65.0	103	0.75%	0.68%	0.024
20	рагелі	65.0		M in ROI:	13809	99,99%	91.36%	2,49%
	DPM Recovered:	15114	DP	M In ROI:	12003	22,22 11	3120010	2 , 42 .2
	DPM Injected:	16580						
	% Recovery:	91.16						

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FIGURE 20

Radiochromatogram of 12-24-hr Pooled Urine from Two Rats after a Single 25-mg/kg Oral Dose of [$^{14}\rm{C}$]Compound I



Sample Name: 01842006.D05

Start: 30-Mar-01

Method: 01842

LSC Reference: LSC966:53-56

. ID: Pooled Urine 12-24hr (X966:17E)

Region No.	¹⁴ C Data Name	Start	Stop	R,	dpm	%ROI	%Total	% Dose
1		6.0	8.0	7.0	466	3.04%	2.82%	0.03%
2	МЗ	8.5	10.5	8.5	6781	57.31%	53.09%	0.64%
4	M9	17.0	19.0	17.0	242	1.58%	1.46%	0.02%
5	MIO	20.0	21.0	20.0	108	0.70%	0.65%	0.01%
7		24.5	25.5	24.5	1370	8.94%	8.28%	0.10%
8	M15	35.5	37.0	35_5	127	0_B35₽	0.77%	0.01%
9	M16	40.0	40.5	40.0	91	0.59%	0.55%	0.01%
10	M17	42.5	44.0	43.5	161	1.05%	0.97%	0.01%
11	M18	44.5	45.5	45.0	157	1.02%	0.95%	0.01%
12	M19	47.5	50.0	48.0	558	3.64%	3,37%	0.04%
14		51.0	52.0	51.5	456	2.9B%	2.76%	0.03%
15 & 16	M20 + M21	52.5	54.0	53.5	1431	9.34%	8.65%	0.10%
17	M23	56.0	57.0	56.5	794	5.18%	4.80%	0.06%
18	M24	59.5	61.0	60.0	204	1.33%	1.23%	0.01%
	14154	63.5	64.5	64.0	196	1.28%	1.19%	0.01%
19	nomel	65.0	65.5	65.0	1B1	1.18%	1.09%	0.01%
20	DPM Recovered:	16540		in ROI:	15323	99.99%	92.63%	1.10%

DPM Injected: 16975

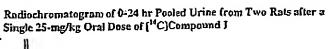
% Recovery: 97.44

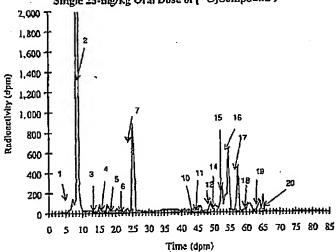
WQ 2004/037211

PCT/US2003/034083

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FIGURE 21





Sample Name: 01842008.D02

Method: 01842

1D: X966:28A (Pooled Urins, 0-24 hr)

Start: 3-Apr-01 LSC Reference: LSC966:76-79

Region	™C Data						•	
Np.	Name	Start	Stop	R,	dpm	%ROI	%7௦௰	% Dose
		6.0	7.5	7.0	306	2.02%	1.91%	0.07%
2	мз	0.8	13.0	· 8.5	8998	59.49 %	56.18%	2.19%
3	M7 + M8 + MBA	15.0	16.0	15.0	120	0.79%	0.75%	0.03%
4	M9	17.0	19.5	17.5	238	1.57%	1.49%	0.06%
5	MIO	20.0	22.5	20.5	145	0.96%	0.91%	0.04%
6	M12	23.0	24.0	23.5	93	0.61%	0.58%	0.02%
7	MI3	25.0	26.0	25.0	1409	9,32%	8.80%	0.34%
10	M17	42.0	45.0	44.0	229	1.51%	1.43%	0.06%
11	1410	45.5	46.0	46.0	118	0.78%	0.74%	0.03%
12	. M19	48.5	49.5	49.0	201	1.33%	1.25%	0.05%
14	10112	50.0	51.5	2.0 5	209	1.38%	1.31%	0.05%
	M20	52.0	53.0	51.5	476	3.15%	2,97%	0.12%
15	M21	53.5	56.0	54.5	1197	7.91%	7.47%	0.29%
16		56.5	58.0	57.5	629	4.16%	3.93%	0.15%
17	M23		62.5	61.0	356	2.35%	2,22%	0.09%
18	M24	59.0	64.5	64.0	331	1.53%	1.44%	0.06%
19		63.0	-	65.0	169	1.12%	1.06%	0.04%
20	parent	65.0	65.0			99.98%	94,45%	3.69%
	DPM Recovered:	16015		DPM in ROI:	15124	37.70%	34 A370	2.0,2

DPM Injected: 17175 % Recovery: 93.25